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L1 27251 SEA FILE=REGISTRY ABB=ON PLU=ON PEROXIDE OR PERIODATE OR DIISOCYANATE OR HALOGEN OR BROMOSUCCINIMIDE OR PERMANGANATE OR OZONE OR CHROMIC ACID?/CN OR SULFURYL CHLORIDE?/CN OR SULFOXIDE OR SELENOXIDE
L2 583 SEA FILE=REGISTRY ABB=ON PLU=ON GLUTARALDEHYDE OR DENACOL
L7 786415 SEA FILE=HCAPLUS ABB=ON PLU=ON L1 OR PEROXIDE OR PERIODATE OR DIISOCYANATE OR HALOGEN OR BROMOSUCCINIMIDE OR PERMANGANATE OR OZONE OR CHROMIC(W) ACID? OR SULFURYL(W)CHLORIDE? OR SULFOXIDE OR SELENOXIDE
L8 23988 SEA FILE=HCAPLUS ABB=ON PLU=ON L2 OR GLUTARALDEHYDE OR DENACOL
L9 2683 SEA FILE=HCAPLUS ABB=ON PLU=ON L7 AND L8
L10 24149 SEA FILE=HCAPLUS ABB=ON PLU=ON L7(L)OXIDIZ?
L11 77 SEA FILE=HCAPLUS ABB=ON PLU=ON L10 AND L9
L12 6 SEA FILE=HCAPLUS ABB=ON PLU=ON L11 AND TISSUE

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L12 ANSWER 1 OF 6 HCAPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 2000:772416 HCAPLUS
 DOCUMENT NUMBER: 133:325693
 TITLE: Stabilization of implantable bioprosthetic devices
 INVENTOR(S): Levy, Robert J.; Vyavahare, Narendra
 PATENT ASSIGNEE(S): The Children's Hospital of Philadelphia, USA
 SOURCE: PCT Int. Appl., 20 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000064371	A1	20001102	WO 2000-US11289	20000426
W: CA				
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				

PRIORITY APPLN. INFO.: US 1999-131257 P 19990427
 AB The invention relates to methods of stabilizing glycosaminoglycans in a biol. **tissue** (e.g. a bioprosthetic implant) in conjunction with crosslinking of protein in the **tissue**. The methods of the invention improve the mech. integrity of the device and improves its stability in vivo. The invention also includes biol. **tissues** having stabilized glycosaminoglycans and crosslinked proteins and kits for prep. such **tissues**. An example is given of crosslinking of type I collagen and radiolabeled hyaluronic acid.
 IT 15056-35-6, Periodate
 RL: PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (oxidizing agent; stabilization of implantable bioprosthetic devices)
 IT 111-30-8, Glutaraldehyde
 RL: PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (stabilization of implantable bioprosthetic devices)
 REFERENCE COUNT: 3
 REFERENCE(S):
 (1) Stone; US 4880429 A 1989
 (2) Tardy; US 4931546 A 1990 HCAPLUS
 (3) Trescony; US 5919472 A 1999

L12 ANSWER 2 OF 6 HCAPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1998:183806 HCAPLUS
 DOCUMENT NUMBER: 128:275021
 TITLE: Chemical modification of titanium surfaces for covalent attachment of biological molecules
 AUTHOR(S): Nanci, A.; Wuest, J. D.; Peru, L.; Brunet, P.; Sharma, V.; Zalzal, S.; McKee, M. D.
 CORPORATE SOURCE: Faculty of Dentistry, Universite de Montreal, Montreal, PQ, H3C 3J7, Can.
 SOURCE: J. Biomed. Mater. Res. (1998), 40(2), 324-335
 CODEN: JBMRBG; ISSN: 0021-9304
 PUBLISHER: John Wiley & Sons, Inc.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB The surface of implantable biomaterials is in direct contact with the host **tissue** and plays a crit. role in detg. biocompatibility. In order

to improve the integration of implants, it is desirable to control interfacial reactions such that nonspecific adsorption of proteins is minimized and **tissue**-healing phenomena can be controlled. In this regard, our goal has been to develop a method to functionalize **oxidized** titanium surfaces by the covalent immobilization of bioactive org. mols. Titanium first was chem. treated with a mixt. of sulfuric acid and hydrogen **peroxide** to eliminate surface contaminants and to produce a consistent and reproducible titanium oxide surface layer. An intermediary aminoalkylsilane spacer mol. was then covalently linked to the oxide layer, followed by the covalent binding of either alk. phosphatase or albumin to the free terminal NH₂ groups using **glutaraldehyde** as a coupling agent. Surface analyses following coating procedures consisted of XPS, SEM, and at. force microscopy (AFM). Enzymic activity of coupled alk. phosphatase was assayed colorimetrically, and surface coverage by bound albumin was evaluated by SEM visualization of colloidal gold immunolabeling. Our results indicate that the linkage of the aminoalkylsilane to the **oxidized** surface is stable and that bound proteins such as alk. phosphatase and albumin retain their enzymic activity and antigenicity, resp. The d. of immunolabeling for albumin suggests that the binding and surface coverage obtained is in excess of what would be expected for inducing biol. activity. In conclusion, this method offers the possibility of covalently linking selected mols. with known biol. activity to **oxidized** titanium surfaces in order to guide and promote the **tissue** healing that occurs during implant integration in bone and soft **tissues**.

- IT 13463-67-7D, Titanium oxide, conjugates with aminoalkylsilane and alk. phosphatase
 RL: DEV (Device component use); PEP (Physical, engineering or chemical process); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (chem. modification of titanium surfaces for covalent attachment of biol. mols.)
- IT 7722-84-1, Hydrogen **peroxide**, reactions
 RL: RCT (Reactant)
 (chem. modification of titanium surfaces for covalent attachment of biol. mols.)

- L12 ANSWER 3 OF 6 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1997:384915 HCPLUS
 DOCUMENT NUMBER: 127:67604
 TITLE: Tanning effect of dialdehydestarch and dialdehydedextrin
 AUTHOR(S): Sugiyama, Rika; Chonan, Yasumasa; Okamura, Hiroshi
 CORPORATE SOURCE: Japan
 SOURCE: Hikaku Kagaku (Sci.) (1997), 43(1), 55-63
 CODEN: HIKAEJ; ISSN: 0018-1811
 PUBLISHER: Nippon Hikaku Gijutsu Kyokai
 DOCUMENT TYPE: Journal
 LANGUAGE: Japanese
 AB The tanning effect of dialdehydestarch was compared with that of formaldehyde and **glutaraldehyde**, which are generally used as aldehyde tanning agents. The tanning effect of dialdehydedextrin, which is an oxide of dextrin (an intermediate product in the hydrolysis of starch) prep'd. by metaperiodate, was also studied. The results were as follows: (1) Dextrin from potato starch was **oxidized** by metaperiodate. After an excess of the **periodate** was removed by the use of ethylene glycol, the product was dialyzed by distd. water, concd. under reduced pressure and freeze-dried, so that a dialdehydedextrin prep'd. with an oxidn. rate (aldehyde content) of 87% was

produced. (2) Dialdehydedextrin was found to more quickly penetrate into hide tissue than dialdehydestarch and to have tanning ability.

IT 9047-50-1P, Dialdehydestarch
 RL: PRP (Properties); SPN (Synthetic preparation); TEM (Technical or engineered material use); PREP (Preparation); USES (Uses)
 (prep. and tanning effect of dialdehydestarch and dialdehydedextrin)

L12 ANSWER 4 OF 6 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1983:68248 HCPLUS
 DOCUMENT NUMBER: 98:68248
 TITLE: The cytochemical reactivity of cerium ions with cardiac muscle
 AUTHOR(S): Christie, Keith N.; Stoward, Peter J.
 CORPORATE SOURCE: Dep. Anat., Univ. Dundee, Dundee, UK
 SOURCE: Acta Histochem. Cytochem. (1982), 15(5), 656-72
 CODEN: ACHCBO
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB When unfixed cardiac muscle from the rat or hamster was incubated in substrate-free media contg. CeCl₃ buffered to pH 7.5 with Tris-maleate-sucrose, electron-dense reaction bodies were formed in the sarcoplasm and, to a greater extent, inside and budding from mitochondria. Fixation in glutaraldehyde prevented the reaction product appearing within mitochondria, but had no effect on either the bodies assocd. with outer mitochondrial membranes or those apparently free in the sarcoplasm or outside the cells. No reaction product was obsd. in unfixed liver, kidney, or skeletal muscle. The electron-dense reaction product within cardiocytes was absent in fixed tissues treated beforehand with acetone and in fresh tissues preincubated with the D-amino acid oxidase inhibitor kojic acid. It was substantially reduced by preincubation in Tiron (a superoxide scavenger), diethyldithiocarbamic acid (an inhibitor of Cu-contg. enzymes such as diamine oxidase and superoxide dismutase), or EDTA, and totally if any of these compds. was included in the incubation medium. However, p-chloromercuribenzene-sulfonate, a nonpenetrating and nonspecific inhibitor of NAD(P)H oxidoreductases, and atebrine limited the amt. of reaction product formed in all sites by much less. Chlorpromazine inhibited the formation of reaction product in the sarcoplasm but gave rise to a coagulated product within mitochondria. With clorgyline, in contrast, no product was deposited in mitochondria and large amorphous masses of reaction product appeared in the sarcoplasm. None of the following, when included in the incubation medium, had any effect: 3-amino-1,2,4-triazole, catalase, dicoumarol, GSH peroxidase, mannitol, MeOH, Pargyline, o-phenanthroline, KCN, rotenone, NaN₃, Na benzoate, Na pyruvate, superoxide dismutase, and thiourea. Evidently, the electron-dense product arises from a reaction of ions with either a lipid peroxide or endogenous H₂O₂ generated by a metal-contg. thiol enzyme, possibly an oxidized form of monoamine oxidase A with the properties of a diamine oxidase. The significance of the Ce reaction in the ageing of cells is discussed briefly.

L12 ANSWER 5 OF 6 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1981:26897 HCPLUS
 DOCUMENT NUMBER: 94:26897
 TITLE: Radioimmunoassay of methionine5-enkephalin sulfoxide: phylogenetic and anatomical distribution
 AUTHOR(S): King, Judy A.; Millar, Robert P.
 CORPORATE SOURCE: Med. Sch., Univ. Cape Town, Observatory, 7925, S. Afr.

SOURCE: Peptides (Fayetteville, N. Y.) (1980), 1(3), 211-16
 CODEN: PPTDD5; ISSN: 0196-9781

DOCUMENT TYPE: Journal
 LANGUAGE: English

AB A sensitive and specific radioimmunoassay (RIA) for the **oxidized** form of methionine5-enkephalin (Met5-Enk), met5-Enk **sulfoxide** (Met5-Enk-S), was developed. Antisera were raised in rabbits against Met5-Enk coupled to carrier proteins with **glutaraldehyde** or carbodiimide. Displacement of ¹²⁵I-labeled Met5-Enk bound to antiserum by Met5-Enk was poor, but Met5-Enk-S displayed good displacement suggesting that the Met5-Enk immunogen was **oxidized** to Met5-Enk-S and that the antisera were formed against this compd. The sensitivity of the RIA for Met5-Enk-S was 0.02 pmole/tube using the most sensitive antiserum. The antisera showed negligible cross-reactivity with leucine5-enkephalin and with both native and **oxidized** endorphins. Cross-reactivity was 15-28% with the fragment Met5-Enk (2-5) **sulfoxide** and 9-25% with D-Ala2-Met5-Enk **sulfoxide**. The antisera showed <0.01% cross-reactivity with other Met5-Enk fragments and naturally occurring neuropeptides. **Tissue** exts. were **oxidized** with H₂O₂ prior to assay. Met5-Enk-S immunoreactivity (IMR) was detected in brain, pituitary gland, pancreas, and intestine exts. of the rat, chicken, toad (*Xenopus laevis*) and teleost (*Sarotherodon mossambicus*), and in cerebral-subesophageal ganglion exts. of the snail (*Helix*). All **tissue** exts. showed parallelism in serial diln. to synthetic mammalian Met5-Enk-S, suggesting possible immunol. identity. Thus, spontaneous oxidn. of Met5-Enk immunogen occurs such that antisera are produced against the **sulfoxide** analog of Met5-Enk, and may account for the relative insensitivity of some published RIAs using Met5-Enk std. These findings demonstrate a wide phylogenetic and anatomical distribution of Met5-Enk IMR.

IT 60283-51-4
 RL: ANT (Analyte); ANST (Analytical study)
 (detn. of, in animal **tissues** by radioimmunoassay, anatomical and phylogenetic distribution in)

L12 ANSWER 6 OF 6 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1975:28168 HCPLUS
 DOCUMENT NUMBER: 82:28168
 TITLE: **Periodate**-lysine-paraformaldehyde fixative.
 New fixative for immunolectron microscopy
 McLean, Ian W.; Nakane, Paul K.
 AUTHOR(S):
 CORPORATE SOURCE: Med. Cent., Univ. Colorado, Denver, Colo., USA
 SOURCE: J. Histochem. Cytochem. (1974), 22(12), 1077-83
 CODEN: JHCYAS
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB A new fixative that primarily stabilizes carbohydrate moieties was developed for immunolectron microscopy. It contained **periodate**, lysine, and paraformaldehyde. Theor., the carbohydrates are **oxidized** by **periodate** and cross-linked by lysine. The fixative can preserve antigenicity as well as paraformaldehyde and ultrastructure as well as **glutaraldehyde**. By using this fixative and the peroxidase-labeled antibody technique, basement membrane antigen was localized within the cisternae of endoplasmic reticulum of parietal yolk-sac cells and in extracellular basement membranes with adequate **tissue** preservation, a task that has not been successfully accomplished by conventional fixatives.

IT 7790-28-5
 RL: USES (Uses)

(in fixative, for carbohydrates stabilization)

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L1 27251 SEA FILE=REGISTRY ABB=ON PLU=ON PEROXIDE OR PERIODATE OR DIISOCYANATE OR HALOGEN OR BROMOSUCCINIMIDE OR PERMANGANATE OR OZONE OR CHROMIC ACID?/CN OR SULFURYL CHLORIDE?/CN OR SULFOXIDE OR SELENOXIDE

L2 583 SEA FILE=REGISTRY ABB=ON PLU=ON GLUTARALDEHYDE OR DENACOL

L7 786415 SEA FILE=HCAPLUS ABB=ON PLU=ON L1 OR PEROXIDE OR PERIODATE OR DIISOCYANATE OR HALOGEN OR BROMOSUCCINIMIDE OR PERMANGANATE OR OZONE OR CHROMIC(W) ACID? OR SULFURYL(W)CHLORIDE? OR SULFOXIDE OR SELENOXIDE

L8 23988 SEA FILE=HCAPLUS ABB=ON PLU=ON L2 OR GLUTARALDEHYDE OR DENACOL

L9 2683 SEA FILE=HCAPLUS ABB=ON PLU=ON L7 AND L8

L10 24149 SEA FILE=HCAPLUS ABB=ON PLU=ON L7(L)OXIDIZ?

L11 77 SEA FILE=HCAPLUS ABB=ON PLU=ON L10 AND L9

L12 6 SEA FILE=HCAPLUS ABB=ON PLU=ON L11 AND TISSUE

L13 213 SEA FILE=HCAPLUS ABB=ON PLU=ON (L9 AND ?TISSUE?) NOT L12

L20 23 SEA FILE=HCAPLUS ABB=ON PLU=ON L13 AND (HEAT OR ?RADIAT? OR UV OR ULTRAVIOL? OR GAMMA)

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L20 ANSWER 1 OF 23 HCAPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 2001:850911 HCAPLUS
 DOCUMENT NUMBER: 135:376709
 TITLE: Protein matrix materials, devices and methods of
 making and using thereof
 INVENTOR(S): Masters, David B.
 PATENT ASSIGNEE(S): Gel-Del Technologies, Inc., USA
 SOURCE: PCT Int. Appl., 158 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001087267	A1	20011122	WO 2001-US6502	20010228
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.:			US 2000-185420	P 20000228
			US 2000-222762	P 20000803

AB The present invention relates to protein matrix materials and devices and the methods of making and using protein matrix materials and devices. More specifically the present invention relates to protein matrix materials and devices that may be utilized for various medical applications including, but not limited to, drug delivery devices for the

controlled release of pharmacol. active agents, encapsulated or coated stent devices, vessels, tubular grafts, vascular grafts, wound healing devices including protein matrix suture material and meshes, skin/bone/tissue grafts, biocompatible electricity conducting matrixes, clear protein matrixes, protein matrix adhesion prevention barriers, cell scaffolding and other biocompatible protein matrix devices. Furthermore, the present invention relates to protein matrix materials and devices made by forming a film comprising one or more biodegradable protein materials, one or more biocompatible solvents and optionally one or more pharmacol. active agents. The film is then partially dried, rolled or otherwise shaped, and then compressed to form the desired protein matrix device.

IT 111-30-8, Glutaraldehyde

RL: RCT (Reactant)

(crosslinking agent; protein matrix materials, devices and methods of making and using thereof)

IT 9054-89-1, Superoxide dismutase

RL: PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
(protein matrix materials, devices and methods of making and using thereof)

IT 67-68-5, Dmso, uses

RL: NUU (Other use, unclassified); USES (Uses)

(solvent; protein matrix materials, devices and methods of making and using thereof)

REFERENCE COUNT:

1

REFERENCE(S): (1) Biegajaski; US 5700478 A 1997 HCPLUS

L20 ANSWER 2 OF 23 HCPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:697421 HCPLUS

DOCUMENT NUMBER: 135:315588

TITLE: Magnetosensitive microcarrier for culturing animal cells and its preparing process

INVENTOR(S): Wei, Cong; Lu, Xiuju; Ouyang, Fan

PATENT ASSIGNEE(S): Inst. of Chemical Metallurgy, Chinese Academy of Sciences, Peop. Rep. China

SOURCE: Faming Zhuanli Shengqing Gongkai Shuomingshu, 12 pp.

CODEN: CNXXEV

DOCUMENT TYPE: Patent

LANGUAGE: Chinese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
CN 1289842	A	20010404	CN 1999-119643	19990923

AB The microcarrier with size of 120-300 .mu.m has core-shell structure. The magnetosensitive microsphere as core is prep'd. by silylating 0.1-5 .mu.m magnetic powder with silane in 0.1-1% acetic acid for 10 min, drying at <100.degree., mixing with monomer, crosslinking agent, initiator, and 3-8% gelatin soln., polymg. at 80-90.degree. for 3-4 h, aging at 90-100.degree. for 2 h, washing with water, soaking in ethanol overnight, soaking in **chromic acid** or sulfonic acid for >3 min, and drying. The microcarrier is prep'd. by dissolving gelatin or glucose in phosphate buffer (pH 5.0-7.2) to obtain 5-20% gelatin soln., adding the magnetosensitive microsphere to the soln., stirring for 5 min, dispersing in Sp80 contg. toluene for >5 min, crosslinking with 10-50% **glutaraldehyde** soln. for 1-3 h, dispersing in phosphate buffer (pH 7.0-7.2), regulating with satd. H3BO3 soln. to pH 9-10, reducing with 0.5-2% NaBH4 for 10-24 h, washing with phosphate buffer, drying at

<70.degree., adding 1-3M NaOH soln., stirring for >1 h, washing, and drying at <70.degree. or storing in 70% ethanol. The magnetic powder is .
gamma.-Fe(OH)2, **.gamma.**-Fe2O3, or Fe3O4. The monomer is styrene, 4-methylstyrene, 3-methylstyrene, **.alpha.**-methylstyrene, or Me methacrylate. The crosslinking agent is divinylbenzene or triallyl isocyanurate. The initiator is azodiisobutyronitrile.

- IT 7738-94-5, **Chromic acid** (H2CrO4)
 RL: NUU (Other use, unclassified); USES (Uses)
 (magnetosensitive microcarrier for culturing animal cells and prepg. process)
- IT 111-30-8, **Glutaraldehyde**
 RL: RCT (Reactant)
 (magnetosensitive microcarrier for culturing animal cells and prepg. process)

L20 ANSWER 3 OF 23 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 2000:456909 HCPLUS
 DOCUMENT NUMBER: 133:79314
 TITLE: Method for producing autogenous vaccines for treating chlamydial infections in mammals and humans
 INVENTOR(S): Sonntag, Hans-Guenther; Nolte, Oliver; Weiss, Hannelore; Weiss, Hans-Erich
 PATENT ASSIGNEE(S): Germany
 SOURCE: PCT Int. Appl., 10 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: German
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000038712	A2	20000706	WO 1999-EP9993	19991216
WO 2000038712	A3	20001102		
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
DE 19860438	C1	20000907	DE 1998-19860438	19981228
EP 1140160	A2	20011010	EP 1999-967946	19991216
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
PRIORITY APPLN. INFO.:			DE 1998-19860438 A	19981228
			WO 1999-EP9993	W 19991216

- AB The title vaccine is produced by heating **tissue** material contg. chlamidia to >50.degree., preferably 80-85.degree., for 1-5 h in the presence of protein-crosslinking agents, preferably formaldehyde or paraformaldehyde. Thus, fresh placenta from a chlamidia-infected aborted lamb was heated at 95.degree. for 60 min, minced, sieved, dild. 1:3 with sterile, pyrogen-free physiol. NaCl soln. contg. 0.3 vol.% formalin, heated at 80.degree. for 2 h, and incubated at 37.degree. for 24 h to verify sterility. Sheep inoculated 3-4 times with this vaccine showed a decrease in spontaneous abortion rate from 20-40% to <3%.
- IT 111-30-8, Pentanedral 7790-28-5

RL: RCT (Reactant)
 (protein-crosslinking agent; method for producing autogenous vaccines
 for treating chlamydial infections)

L20 ANSWER 4 OF 23 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1999:746685 HCPLUS
 DOCUMENT NUMBER: 132:134306
 TITLE: A solid-phase assay for quantitative analysis of sulfated glycosaminoglycans at the nanogram level.
 Application to tissue samples
 AUTHOR(S): Vynios, D. H.; Faraos, A.; Spyracopoulou, G.; Aletras, A. J.; Tsiganos, C. P.
 CORPORATE SOURCE: Department of Chemistry, Laboratory of Biochemistry, University of Patras, Patras, 261 10, Greece
 SOURCE: J. Pharm. Biomed. Anal. (1999), 21(4), 859-865
 CODEN: JPBADA; ISSN: 0731-7085
 PUBLISHER: Elsevier Science B.V.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB A sensitive and accurate solid-phase methodol. for the quant. anal. of glycosaminoglycans is described. Chondroitin-4-sulfate (CSA) was labeled with biotin hydrazide after the reaction of its carboxyl groups with it in the presence of carbodiimide. Polystyrene plates modified with sequential reaction with glutaraldehyde (GH) and spermine to possess amino groups were used to immobilize electrostatically the biotin labeled CSA. Exogenously added sulfated glycosaminoglycans (GAGS) [variously sulfated chondroitin sulfates and heparan sulfate (HS)] were found to compete to this immobilization in a concn. dependent mode, within a concn. range from 10 up to 300 ng/mL. Glycosaminoglycan-derived oligosaccharides competed to a degree similar to that of intact mols. Hyaluronan (HA) and keratan sulfate (KS) did not compete the immobilization. The procedure was applied for the rapid and reproducible detn. of the sulfated glycosaminoglycans in proteinase digests of small tissue samples or cell cultures with high sensitivity and accuracy.
 IT 111-30-8, Glutaraldehyde
 RL: ARU (Analytical role, unclassified); ANST (Analytical study)
 (a solid-phase assay for quant. anal. of sulfated glycosaminoglycans at nanogram level, application to tissue samples)
 IT 9003-53-6, Polystyrene
 RL: DEV (Device component use); USES (Uses)
 (a solid-phase assay for quant. anal. of sulfated glycosaminoglycans at nanogram level, application to tissue samples)
 REFERENCE COUNT: 38
 REFERENCE(S):
 (1) Bitter, T; Anal Biochem 1962, V4, P330 HCPLUS
 (2) Bjornsson, S; Anal Biochem 1998, V256, P229
 HCPLUS
 (4) Dische, Z; J Biol Chem 1947, V167, P189 HCPLUS
 (6) Farndale, R; Connect Tissue Res 1982, V9, P247
 HCPLUS
 (7) Fosang, A; Matrix 1990, V10, P306 HCPLUS
 ALL CITATIONS AVAILABLE IN THE RE FORMAT
 L20 ANSWER 5 OF 23 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1999:265982 HCPLUS
 DOCUMENT NUMBER: 130:316630
 TITLE: Medicinal carrier particle for tissue-specific application
 INVENTOR(S): Mueller, Rainer; Lueck, Martin; Kreuter, Joerg
 PATENT ASSIGNEE(S): DDS Drug Delivery Service Gesellschaft zur Foerderung

der Forschung in pharmazeutischer Technologie und
Biopharmazie m.b.H., Germany

SOURCE: Ger. Offen., 18 pp.
CODEN: GWXXBX

DOCUMENT TYPE: Patent
LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 19745950	A1	19990422	DE 1997-19745950	19971017
WO 9920256	A2	19990429	WO 1998-EP6429	19981013
WO 9920256	A3	19990819		
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 9912272	A1	19990510	AU 1999-12272	19981013
EP 1023052	A2	20000802	EP 1998-955425	19981013
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
US 6288040	B1	20010911	US 2000-529600	20000621
PRIORITY APPLN. INFO.:			DE 1997-19745950 A	19971017
			WO 1998-EP6429 W	19981013

- AB Drug carrier particles are provided for delivery of drugs across the blood-brain barrier to the central nervous system for treatment of central nervous disorders. The particles, in drug-loaded or drug-free form, bear on their surface .gtoreq.1 covalently bound or adsorbed recognition protein (e.g. an apolipoprotein) for receptors in the brain or blood-brain barrier, or the particle surface is modified (e.g. with an ethoxylated surfactant) so that a recognition protein is bound on contact with the particle. Thus, poly(Bu cyanoacrylate) nanoparticles loaded with the analgesic, dalargin, were surface modified with Tween 80 and incubated with apolipoprotein E. Administration of these nanoparticles i.v. to mice produced an analgesic effect, as shown in the tail-flick test.
- IT 111-30-8, Glutardialdehyde 7790-21-8, Potassium periodate 7790-28-5, Sodium periodate
RL: RCT (Reactant)
(drug carrier particle surface activation with; medicinal carrier particle for tissue-specific application to brain)

L20 ANSWER 6 OF 23 HCPLUS COPYRIGHT 2001 ACS
ACCESSION NUMBER: 1995:555080 HCPLUS
DOCUMENT NUMBER: 123:3802
TITLE: **Tissue** thromboplastin
AUTHOR(S): Baykeev, R. F.; Azancheev, N. M.; Chernov, A. N.; Filippov, A. V.; Dvorancev, S. N.
CORPORATE SOURCE: Kazan State Medical University, Kazan 420012, Tatarstan, Russia
SOURCE: J. Mol. Struct. (1995), 348, 305-8
DOCUMENT TYPE: Journal
LANGUAGE: English
AB The dynamic structure of **tissue** thromboplastin (**tissue**

factor, TF) from the human brain has been studied. The phase state of lipids, the mobility of mol. segments in the structure of protein and lipid moieties have been detd. to permit us to elaborate the dynamic model of **tissue** thromboplastin from the human brain.

IT 67-68-5, Dimexidum, biological studies 111-30-8,

Glutaraldehyde

RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)
(**tissue** thromboplastin structure response to)

L20 ANSWER 7 OF 23 HCPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1994:676202 HCPLUS

DOCUMENT NUMBER: 121:276202

TITLE: Macromolecular microparticles, methods of production, and diagnostic, therapeutic, and other uses thereof

INVENTOR(S): Woiszwillo, James E.

PATENT ASSIGNEE(S): Middlesex Sciences, Inc., USA

SOURCE: PCT Int. Appl., 74 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 6

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9420856	A1	19940915	WO 1994-US2316	19940304
W: AU, CA, HU, JP, KR, PL RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
CA 2157793	AA	19940915	CA 1994-2157793	19940304
AU 9463585	A1	19940926	AU 1994-63585	19940304
EP 688429	A1	19951227	EP 1994-910826	19940304
EP 688429	B1	19980211		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
JP 08507806	T2	19960820	JP 1994-520171	19940304
US 5578709	A	19961126	US 1994-206456	19940304
EP 809110	A1	19971126	EP 1997-112821	19940304
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE AT 163230	E	19980215	AT 1994-910826	19940304
ES 2113094	T3	19980416	ES 1994-910826	19940304
PRIORITY APPLN. INFO.:			US 1993-28237	A 19930309
			EP 1994-910826	A3 19940304
			WO 1994-US2316	W 19940304

AB A macromol. microparticle compn. is formed by dehydrating an aq. macromol. soln. and crosslinking the dehydrated macromols. with a crosslinking agent while in a liq. phase or with heat. Preferably, the dehydrating agent is a polymer mixt. of polyvinylpyrrolidone and polyethylene glycol, the crosslinking reagent is **glutaraldehyde**, and the macromol. is a protein, most preferably an Ig. Methods of use for research, diagnostics and therapeutics are also provided. Prepn. of e.g. tetanus toxoid microparticles and immunization of mice with the tetanus toxoid microparticles is described.

IT 111-30-8, **Glutaraldehyde**

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(crosslinking agent; macromol. microparticles, methods of prodn., and diagnostic, therapeutic, and other uses thereof)

IT 9003-99-0D, Peroxidase, IgG conjugates

RL: BUU (Biological use, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(macromol. microparticles, methods of prodn., and diagnostic, therapeutic, and other uses thereof)

L20 ANSWER 8 OF 23 HCAPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1993:533233 HCAPLUS
 DOCUMENT NUMBER: 119:133233
 TITLE: The Influence of chemical structure on the extent and sites of carcinogenesis for 522 rodent carcinogens and 55 different human carcinogen exposures
 AUTHOR(S): Ashby, J.; Paton, D.
 CORPORATE SOURCE: Cent. Toxicol. Lab., ICI, Macclesfield/Ches., SK10 4TJ, UK
 SOURCE: Mutat. Res. (1993), 286(1), 3-74
 CODEN: MUREAV; ISSN: 0027-5107
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB L. S. Gold et al. (1991) tabulated the results of rodent bioassays on 522 chems. and analyzed the data. The present study complements those analyses by providing a perspective from the viewpoint of the chem. structure of the carcinogens. The chem. structure of each of the carcinogens is displayed and the Gold database is represented with the test agents as the primary variable. The carcinogens are gathered into 6 chem. classes and each chem. is assessed for structural alerts to DNA reactivity. The database is then analyzed using an integration of the following parameters: bioassay in rat, mouse or both; structural alert status; chem. class; sites and multiplicity of carcinogenesis, and trans-species carcinogenicity. A series of figures is presented that enables rapid acquaintance with what represents the core database of rodent carcinogenicity. The several analyses presented combine in endorsing the reality of two broad classes of rodent carcinogen, presumed DNA-reactive and others (putative genotoxic and non-genotoxic carcinogens, but semantics have been largely avoided). H. M. Vainio et al. (1991) and his colleagues have tabulated 55 situations in which humans have succumbed to chem. induced cancer and have listed the **tissues** affected. This database of human carcinogens has been analyzed in the present study as done for the rodent carcinogen database, and comparisons made between the two. The predominance of putative genotoxic carcinogens in the human database was confirmed, as was the reality of putative non-genotoxic carcinogenicity in humans. It is concluded that putative genotoxic rodent carcinogenesis can be correlated both with chem. structure and the extent and nature of the induced effect, and that it is of clear relevance to humans. In contrast, it is concluded that putative non-genotoxic rodent carcinogenesis is more closely related to the test species than to the test chem., and that it is essentially unpredictable in the absence of mechanistic models.
 IT 91-93-0 120-62-7, Piperonyl sulfoxide
 122-60-1 7722-84-1, Hydrogen peroxide,
 biological studies
 RL: ADV (Adverse effect, including toxicity); PRP (Properties); BIOL (Biological study)
 (neoplasm from, of **tissues**, in lab. animals, structure role in, human in relation to)
 L20 ANSWER 9 OF 23 HCAPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1993:490604 HCAPLUS
 DOCUMENT NUMBER: 119:90604
 TITLE: Freeze-substitution fixation for immunohistochemistry at the light microscopic level: effects of solvent and chemical fixatives

AUTHOR(S): Yamashita, Shuji; Yasuda, Kenjiro
 CORPORATE SOURCE: Keio J. Coll. Nurs., Tokyo, 160, Japan
 SOURCE: Acta Histochem. Cytochem. (1992), 25(6), 641-50
 CODEN: ACHCBO; ISSN: 0044-5991

DOCUMENT TYPE: Journal
 LANGUAGE: English

AB The suitability of the freeze-substitution and paraffin embedding technique for the immunolocalization of membrane-bound and sol. antigens in the rat at the light microscopic level was investigated. Frozen **tissues** were substituted at -80.degree. in methanol- or acetone-contg. chem. fixatives, i.e., formaldehyde, **glutaraldehyde**, acrolein, hexamethylene **diisocyanate** (HMD), di-Et malonimide (DEM) and 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide (water-sol. carbodiimide, WSC). The following antigens were localized using monoclonal antibodies and antisera: (A) membrane-bound antigens; **gamma**.-glutamyl transpeptidase (**gamma**.-GTP), common antigen of secretory granule membrane (SG 170 antigen) and Golgi-assocd. antigen (GAA 108); and (B) sol. antigens; branched-chain amino acid transferase type I isoenzyme (BAT), glutamate dehydrogenase (GDH), pancreatic amylase, proliferating cell nuclear antigen (PCNA), rat serum albumin and IgG. (1) The freeze-substitution technique maintained an excellent **tissue** structure and conservation of antigenicity. By using this method, BAT was localized in mitochondria in liver cells, and **gamma**.-GTP was demonstrated in the secretory granule membrane of pancreatic acinar cells, although conventional fixation methods provided neg. reaction. (2) In general, the membrane-bound antigens were localized in detail with a strong immunoreaction in the **tissues** substituted in solvent alone; however, for the localization of sol. antigens, **tissues** substituted in solvents contg. chem. fixatives revealed a strong and precise antigen localization. (3) Formaldehyde and **glutaraldehyde** proved to be better fixatives concerning the conservation of structure and antigenicity than the other chem. reagents. (4) The choice of substitution solvent was important for the immunohistochem. of some antigens. For example, methanol was suitable for PCNA, and substitution in acetone was essential for **gamma**.-GTP using one of the monoclonal antibodies to **gamma**.-GTP.

IT 111-30-8, **Glutaraldehyde** 822-06-0,
Hexamethylene diisocyanate
 RL: BIOL (Biological study)
 (freeze-substitution fixation for immunolocalization of membrane-bound and sol. antigens in relation to)

L20 ANSWER 10 OF 23 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1992:441497 HCPLUS
 DOCUMENT NUMBER: 117:41497
 TITLE: DNA-peroxidase probing of some plant and animal virus infections
 AUTHOR(S): Drygin, Yu. F.; Buzmakov, A. V.; Teterina, N. L.; Morozov, S. Yu.
 CORPORATE SOURCE: A. N. Belozerskii Lab. Mol. Biol. Bioorg. Chem., M. V. Lomonosov Moscow State Univ., Moscow, 119899, Russia
 SOURCE: Mol. Biol. (Moscow) (1992), 26(1), 59-69
 CODEN: MOBIBO; ISSN: 0026-8984
 DOCUMENT TYPE: Journal
 LANGUAGE: Russian
 AB DNA-peroxidase probes were synthesized for the detection of .lambda. phage DNA (model system), polio, potato X (PVX) and M (PVM) tobacco mosaic (TMV) viral RNAs by spot hybridization onto nitrocellulose membranes. The cDNAs (300-1400 bases) complementary to the viral RNAs were cloned in M13 phage

DNA or plasmid pTZ19. Each step of the probe construction and the diagnostic procedure were thoroughly examd. Peroxidase activity obsd. with non-toxic stain (NTS) was 3-5 fold more sensitive than were .alpha.-Cl-naphthol or bisanisidine. Horseradish peroxidase (HRP) became much more stable to **heat** in dild. samples and was 2-3 fold more active after coupling with a polyethylenimine (PEI) spacer. Also, sodium borohydride redn. of the cDNA and PEI-HRP adduct crosslinked by **glutaraldehyde** resulted in the stabilization of the probes. Target nucleic acids or diagnostic samples were efficiently fixed onto nitrocellulose membranes by a short-time UV irradn. Viral detection in cellular exts. with prep'd. probes takes 4-5 h with 100-200 ng/mL of specific nucleotide sequence. Up to 20 pg (<10-17M) of the purified viral nucleic acids or 30-50 pg in the total fraction of the cellular nucleic acids isolated from the infected cells were identified with the probes. A 500-10,000-fold dild. lysate of the HeLa cells infected with poliovirus (PV1) and both crude exts. of potato tuber of potato and tobacco leaf **tissues** infected with PVX, PVM or TMV displayed specific signals with the resp. DNA-HRP probes.

IT 9003-99-0, Peroxidase

RL: USES (Uses)

(horseradish, DNA probes contg., for plant and animal DNA and RNA virus detection by spot hybridization, construction of)

L20 ANSWER 11 OF 23 HCPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1991:577874 HCPLUS

DOCUMENT NUMBER: 115:177874

TITLE: Exact ultrastructural localization of glutathione peroxidase in normal rat hepatocytes: advantages of microwave fixation

AUTHOR(S): Utsunomiya, Hirotoshi; Komatsu, Noriyuki; Yoshimura, Shinichi; Tsutsumi, Yutaka; Watanabe, Keiichi

CORPORATE SOURCE: Sch. Med., Tokai Univ., Isehara, Japan

SOURCE: J. Histochem. Cytochem. (1991), 39(9), 1167-74

CODEN: JHCYAS; ISSN: 0022-1554

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Glutathione peroxidase (GSH-PO), a highly sol., selenium-dependent enzyme metabolizing lipid **peroxides**, is allegedly distributed in both the cytosol and mitochondria. With the pre-embedding method of immunoelectron microscopy for GSH-PO employing conventional immersion-fixation, the nucleic of rat hepatocytes stain pos., whereas mitochondria are neg. Such observations are inconsistent with the results of biochem. and immunoblot analyses using isolated subcellular fractions. In the present study, the combination of microwave irradn. and fixation in 4% paraformaldehyde (PFA), with or without 0.1% **glutaraldehyde** (GA), was used to enhance the accuracy of ultrastructural localization of GSH-PO in rat liver. A small block of liver was **irradiated** by microwave for 10 s in cold cacodylate-buffered 4% PFA contg. 0.1% GA. After further immersion of the **tissue** in 4% PFA at 4.degree. for 1-6 h, the std. procedure for pre-embedding immunoelectron microscopy was employed. Partial inhibition was obsd. of artifactual diffusion of cytosolic GSH-PO into the nuclei; consistent GSH-PO localization in mitochondria was seen. Dual localization of this enzyme in the cytosol and mitochondria of normal rat hepatocytes was thus confirmed.

L20 ANSWER 12 OF 23 HCPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1991:422201 HCPLUS

DOCUMENT NUMBER: 115:22201

TITLE: Superoxide dismutase-catalase conjugates as

INVENTOR(S): **tissue-specific therapeutics**
 Poznansky, Mark J.; Mao, Guo Dong
 PATENT ASSIGNEE(S): University of Alberta, Can.
 SOURCE: PCT Int. Appl., 30 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9103548	A1	19910321	WO 1990-CA279	19900830
W: AU, CA, JP, US				
RW: AT, BE, CH, DE, DK, ES, FR, GB, IT, LU, NL, SE				
CA 2065430	AA	19910301	CA 1990-2065430	19900830
AU 9062854	A1	19910408	AU 1990-62854	19900830
US 5336493	A	19940809	US 1992-836274	19920302
PRIORITY APPLN. INFO.:			GB 1989-19661	19890831
			WO 1990-CA279	19900830

- AB A novel multicomponent conjugate having superoxide dismutase (SOD), catalase, and optionally albumin and a targeting agent such as antibody is provided. A pharmaceutical compn. contg. such conjugate can be used for **tissue-specific** scavenging of superoxide and hydroxyl radicals with higher efficiency than SOD or catalase alone. The half-life of the SOD-catalase conjugates was 300 min in rats. Scavenging of the free radicals using the conjugates was also demonstrated in vitro and in the rat heart model of ischemia-reperfusion.
- IT 9054-89-1DP, Superoxide dismutase, conjugates with catalase
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of, for superoxide and hydroxyl free radical scavenging)
- IT 111-30-8, Glutaraldehyde
 RL: BIOL (Biological study)
 (superoxide dismutase-catalase conjugate prepn. with)

L20 ANSWER 13 OF 23 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1989:131694 HCPLUS
 DOCUMENT NUMBER: 110:131694

TITLE: Use of light microscopic immunotechniques in selecting preparation conditions and immunoprobes for ultrastructural immunolabeling of lactoferrin
 AUTHOR(S): Mutasa, H. C. F.; Pearson, E. C.
 CORPORATE SOURCE: Dep. Haematol. Med., Univ. Cambridge, Cambridge, CB2 2QL, UK
 SOURCE: Histochem. J. (1988), 20(10), 558-66
 CODEN: HISJAE; ISSN: 0018-2214

DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Successful postembedding immunol. labeling for electron microscopy is sometimes difficult to achieve. Light microscopy can be used (1) to detect quickly processing steps that have an adverse effect on the **tissue** antigenicity and (2) to check the specific reactivity of the immunogold detecting system normally employed at the ultrastructural level. The individual steps of fixation, dehydration, and embedding were tested for their ability to preserve antigenicity by light microscopic peroxidase-antiperoxidase cytochem. Steps that severely reduced antigenicity were replaced by less destructive alternatives compatible with reasonable ultrastructural preservation. The specific reactivity of the immunogold detecting system was assessed by using the light

microscopic immunogold Ag staining method. The antigen lactoferrin was studied in human neutrophilic granulocytes from patients with chronic myeloid leukemia. Strong immunolabeling of specific granules and good ultrastructural preservation were obtained by using routine methods at room temp. For lactoferrin, the method of choice was to fix in 3% paraformaldehyde/0.1% glutaraldehyde followed by 1% OsO₄, dehydrate in 70% EtOH, embed in LR White resin, and polymerize at 40.degree. for 40 h. These conditions may not be suitable for all antigens, and for each new antigen a similar study should be carried out.

IT 111-30-8, Pentanodial 15056-35-6, Periodate

RL: BIOL (Biological study)

(lactoferrin antigenicity response to, ultrastructure in relation to)

L20 ANSWER 14 OF 23 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1988:535012 HCAPLUS

DOCUMENT NUMBER: 109:135012

TITLE: Crosslinked peritoneal tissues as novel biomaterials for medical devices and process for their manufacture

INVENTOR(S): Lauren, Mark D.

PATENT ASSIGNEE(S): Australia

SOURCE: U.S., 6 pp.

CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 4755593	A	19880705	US 1986-888717	19860724
AU 8660596	A1	19870129	AU 1986-60596	19850724
PRIORITY APPLN. INFO.:		AU 1985-1616		19850724

AB A biomaterial, suitable for use in medical devices, comprises peritoneum tissue which has been chem. treated to crosslink the collagen in the tissue, rendering the tissue more stable, less antigenic, and sterile. Peritoneum tissue was dissected from the abdominal cavity of calves, the tissue cleaned in phosphate buffered saline, pinned to a polyethylene surface, and exposed to 1% glutaraldehyde in phosphate buffered saline for 24 h at room temp., followed by 2% H₂O₂ for 30 min, and stored in 50% aq. EtOH. The treated tissue had shrinkage temp. 83.5.degree., vs. 66.5 and 67.5.degree. for untreated tissue.

IT 111-30-8, Glutaraldehyde 28605-81-4,

Dicyclohexylmethanedisocyanate

RL: BIOL (Biological study)

(tanning agent for peritoneum tissue, in prepn. of biomaterial)

L20 ANSWER 15 OF 23 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1987:561746 HCAPLUS

DOCUMENT NUMBER: 107:161746

TITLE: Treatment of collagenous tissue for use as xenograft bioprostheses

INVENTOR(S): Milthorpe, Bruce; Schindhelm, Klaus

PATENT ASSIGNEE(S): Domedica Pty. Ltd., Australia

SOURCE: PCT Int. Appl., 16 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 8702880	A1	19870521	WO 1986-AU346	19861113
W: AU, JP, US RW: AT, BE, CH, DE, FR, GB, IT, LU, NL, SE				
AU 8666261	A1	19870602	AU 1986-66261	19861113
AU 586718	B2	19890720		
EP 245383	A1	19871119	EP 1986-906721	19861113
R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE JP 63501344	T2	19880526	JP 1986-506010	19861113
PRIORITY APPLN. INFO.:			AU 1985-3384	19851113
			AU 1984-33	19851113
			WO 1986-AU346	19861113

- AB Collagen-based materials are pretreated with a reagent capable of reacting with or removing the primary amines of the amino acid side chains and then reacted with a crosslinking agent to link some of the remaining amino acid side chains together to prep. implantable prosthetics. Kangaroo tail tendons stored in cold sterile saline soln. were removed and placed in a bicarbonate buffer contg. fluorescein isothiocyanate for 3 day at 4.degree., followed by fixing them in buffered 2% **glutaraldehyde** for 3 day at room temp. The tendons were aseptically transferred to final packaging prior to terminal sterilization by .gamma.-radiation. The prep'd. xenografts were applicable in the treatment of sports injuries of the knee particularly in cruciate ligament repair.
- IT 111-30-8, **Glutaraldehyde** 9047-50-1, Dialdehyde starch
RL: RCT (Reactant)
(crosslinking by, of modified collagens in xenograft prepn.)

L20 ANSWER 16 OF 23 HCPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1987:83050 HCPLUS
 DOCUMENT NUMBER: 106:83050
 TITLE: Cell culture growth enhancement by modified collagens
 INVENTOR(S): Yoshizato, Katsutoshi; Taira, Toshio; Miyata, Teruo
 PATENT ASSIGNEE(S): Koken Co., Ltd., Japan
 SOURCE: Jpn. Kokai Tokkyo Koho, 4 pp.
 CODEN: JKXXAF

DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 61207340	A2	19860913	JP 1985-48350	19850313
JP 05086185	B4	19931210		

- AB Collagens modified by treatment with UV, .alpha.-ray, ethylene oxide, hexamethylene diisocyanate, or glutaryldehyde markedly increase the cell multiplication rate in animal cell culture. Thus, 0.02% sterilized pyrogen-free collagen of calf skin origin was aseptically applied on the surface of a cell culture dish, air-dried, and exposed to 0.5 Mrad .gamma.-ray. When the treated dish was used for cultivation of cell lines of human skin fibroblast, rat liver, and rat epiderm, a marked increase in cell multiplication and growth was obsd.
- IT 111-30-8, **Glutaraldehyde**

RL: BIOL (Biological study)
 (collagen modification by, as animal cell culture growth enhancer)
 IT 822-06-0, Hexamethylenediisocyanate
 RL: BIOL (Biological study)
 (collagen modified by, for cell culture enhancement)

L20 ANSWER 17 OF 23 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1985:450613 HCPLUS
 DOCUMENT NUMBER: 103:50613
 TITLE: Immunochemical localization of
 .alpha.-actinin on Lowicryl-embedded thin-sectioned
tissues
 AUTHOR(S): Lemanski, Larry F.; Paulson, Daniel J.; Hill, Craig
 S.; Davis, Lynn A.; Riles, Linda C.; Lim, Soo Siang
 CORPORATE SOURCE: Upstate Med. Cent., State Univ. New York, Syracuse,
 NY, USA
 SOURCE: J. Histochem. Cytochem. (1985), 33(6), 515-22
 CODEN: JHCYAS; ISSN: 0022-1554
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB A procedure has been developed for the immunochemical localization of intracellular antigens on thin-sectioned **tissues**. The **tissues** were fixed in a **periodate**-lysine-paraformaldehyde soln. or a formaldehyde-**glutaraldehyde** combination and embedded in the acrylate-methacrylate mixt., Lowicryl K4M (Polaron), which was polymd. under UV irradn. at -35.degree.. Thin sections were mounted on gold grids, immunostained using an indirect method with ferritin-labeled antibodies, and, optionally, counterstained with osmium tetroxide and/or lead citrate and uranyl acetate. The procedure provided good morphol. preservation of the cell architecture in adult and embryonic heart, and skeletal and smooth muscle **tissue**, as well as nonmuscle cells. At the same time it retained the antigenicities of several contractile proteins, including myosin, tropomyosin, actin, and .alpha.-actinin. The method has advantages over en bloc staining techniques in that the problem of antibody penetration into the cells is eliminated and careful controls can be performed on adjacent sections. This technique will be useful for localizing, at the ultrastructural level, contractile and other selected proteins in a variety of muscle and nonmuscle cells. Details of the new protocol and a description of the results of using antibody against the contractile protein, .alpha.-actinin, are given.

L20 ANSWER 18 OF 23 HCPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1983:3079 HCPLUS
 DOCUMENT NUMBER: 98:3079
 TITLE: Platelet interaction with the extracellular matrix produced by cultured endothelial cells: a model to study the thrombogenicity of isolated subendothelial basal lamina
 AUTHOR(S): Vlodavsky, I.; Eldor, A.; HyAm, E.; Atzman, R.; Fuks, Z.
 CORPORATE SOURCE: Dep. Radiat., Hadassah Univ. Hosp., Jerusalem, 91120, Israel
 SOURCE: Thromb. Res. (1982), 28(2), 179-91
 CODEN: THBRAA; ISSN: 0049-3848
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Cultured bovine endothelial cells produce an extensive underlying extracellular matrix (ECM) which closely resemble the vascular

subendothelial basal lamina in its organization and chem. compn. This naturally produced ECM was used to study the interaction of platelets with the subendothelium when exposed or covered with vascular endothelial cells. Incubation of platelet-rich plasma with the ECM induced a rapid and massive platelet adherence, aggregation, thromboxane formation, and release reaction. These were demonstrated using phase and SEM, ¹¹¹In- or [¹⁴C]serotonin-labeled platelets, and an RIA for TXB₂. In contrast to the ECM no platelet activation was induced either by uncoated plastic dishes or ECM covered with a confluent endothelial cell monolayer. Aspirinized platelets failed to undergo aggregation and degranulation, when incubated with the ECM. Culture dishes coated with characteristic constituents of the basal lamina such as collagen type IV and type V or fibronectin induced a much lower platelet reactivity as compared with ECM-coated dishes. Digestion of ECM components (collagen, fibronectin, hyaluronic acid, and chondroitin sulfate) by specific enzymes was not assocd. with a substantial decrease in its platelet reactivity. Furthermore, exposure of ECM to SDS or Na **periodate**, or freezing and thawing did not decrease its biol. activity. In contrast, platelet activation was completely abolished following **heat** denaturation or **glutaraldehyde** fixation of the ECM. The availability of a naturally produced ECM provides an appropriate model to study the interaction of platelets with the subendothelium in a controlled system which is isolated from other components of the vessel wall.

L20 ANSWER 19 OF 23 HCPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1980:91814 HCPLUS
 DOCUMENT NUMBER: 92:91814
 TITLE: Cultured sympathetic neurons: effects of cell-derived and synthetic substrata on survival and development
 AUTHOR(S): Hawrot, Edward
 CORPORATE SOURCE: Dep. Neurobiol., Harvard Med. Sch., Boston, MA, 02115, USA
 SOURCE: Dev. Biol. (1980), 74(1), 136-51
 CODEN: DEBIAO; ISSN: 0012-1606
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB Compared to dried collagen films, both 3-dimensional hydrated collagen gels and surfaces coated with basic polymers provided a substratum highly adherent for developing rat sympathetic neurons. Polylysine and polyornithine were suitable for long-term culture when covalently linked with **glutaraldehyde** to an underlying layer of dried gelatin. Dissocd. neurons also attached strongly to a substratum of killed nonneuronal cells fixed by paraformaldehyde, **heat**, EtOH, or TCA. In addn., an extracellular, substrate-assocd. material apparently produced by nonneuronal cells (rat cardiac myocytes and assocd. fibroblasts) promoted the long-term adhesion of growing neurites. The adhesive property of this microexudate was sensitive to trypsin, **periodate**, and alkali, but resistant to hyaluronidase, chondroitinase, 8M urea, and 0.5 HOAc. The choice and development of neurotransmitter function were unaffected by the various substrate tested, except that nonneuronal cells fixed with paraformaldehyde caused a significant induction of cholinergic properties similar to that seen with nonneuronal-conditioned medium.

L20 ANSWER 20 OF 23 HCPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1977:417814 HCPLUS
 DOCUMENT NUMBER: 87:17814
 TITLE: New techniques for cytochemical localization of exogenous peroxidase activity with 2,7-fluorenediamine

AUTHOR(S): and 2,5-fluorenediamine as hydrogen donors
 Ohkawa, Kinichi; Antakly, Tony William; Tanaka,
 Satoshi; Sugai, Naonori

CORPORATE SOURCE: Chest Dis. Res. Inst., Kyoto Univ., Kyoto, Japan

SOURCE: Ann. Histochim. (1976), 21(4), 353-63

CODEN: ANHIAG

DOCUMENT TYPE: Journal

LANGUAGE: English

AB I.v. injected horseradish peroxidase activity was cytochem. detected with 2,7-fluorenediamine (I) and 2,5-fluorenediamine (II) as H-donors. Forty min following i.v. injection of horseradish peroxidase, the liver and kidney were removed. Small pieces of **tissues** were fixed immediately for 7-10 h in chilled phosphate-buffered 1.25% **glutaraldehyde** soln. (pH 7.2). In liver, Kupffer cells and sinusoidal endothelial cells showed staining. In kidney, polymorphonuclear leukocytes and urinary epithelial cells showed staining. Staining was of droplet nature in the light microscope and of vesicular pattern in the electron microscope. Erythrocytes showed weak or no staining. In heat-denatured sections of livers and kidneys, only erythrocytes showed weak staining. Incubation media devoid of either H-donor, I or II, or H-acceptor, H₂O₂, gave rise to no pos. staining. In sections of control animals injected only with physiol. saline soln., erythrocytes, Kupffer cells, and polymorphonuclear leukocytes showed staining; sinusoidal endothelial cells and urinary tubular epithelial cells showed no staining.

IT 9003-99-0

RL: BIOL (Biological study)
 (histochem. localization of exogenous, fluoroenediamine in)

L20 ANSWER 21 OF 23 HCPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1976:118989 HCPLUS

DOCUMENT NUMBER: 84:118989

TITLE: Cytochemical localization of endogenous peroxidase activity in renal medullary collecting tubules and papillary mucosa of the rat

AUTHOR(S): Cavallo, Tito

CORPORATE SOURCE: Sch. Med., Univ. Pittsburgh, Pittsburgh, Pa., USA

SOURCE: Lab. Invest. (1976), 34(3), 223-8

CODEN: LAINAW

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Endogenous peroxidase activity, as demonstrated by the technique of R.C. Graham, Jr., and M.J. Karnovsky (1966), was identified in medullary collecting tubule cells and in the cells of renal papillary mucosa of the rat. Peroxidase reactive sites were obsd. in the perinuclear cisterna, endoplasmic reticulum, and cytoplasmic vesicles of such cells. The specificity of the peroxidase reaction was verified by means of chem. inhibitors (NaN₃, KCN, and aminotriazole), denaturation of the enzyme by heat, exclusion or prior oxidn. of substrate (diaminobenzidine), and high concn. of H₂O₂. Prolonged fixation (**glutaraldehyde**) improved cellular detail but diminished or abolished the peroxidase staining. When exogenous H₂O₂ was excluded from the incubating medium, a pos. reaction was obtained, suggesting that H₂O₂ can be endogenously generated. This observation was confirmed by degradn. of **tissue**-formed H₂O₂ with beef liver catalase and by blocking endogenous generation of H₂O₂ with sodium pyruvate. These studies indicate that the reaction product is the result of an enzymic reaction and that the enzyme is most likely a peroxidase. A similar staining reaction was not obsd. in other tubule segments, including cortical collecting tubules. The

significance of this peroxidase activity is discussed in relation to the cellular localization and biosynthesis of renal medullary prostaglandins.

IT 9003-99-0
 RL: BIOL (Biological study)
 (of kidney medullary collecting tubules and papillary mucosa)

L20 ANSWER 22 OF 23 HCAPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1973:416729 HCAPLUS
 DOCUMENT NUMBER: 79:16729
 TITLE: Effect of various preservation methods on the immunogenicity and antigenic composition of xenogenic valve **tissues** of the heart
 AUTHOR(S): Frolova, M. A.; Barbarash, L. S.; Gudkova, R. G.; Karpinskaya, V. M.
 CORPORATE SOURCE: A. N. Bakulev Inst. Cardiovasc. Surg., Moscow, USSR
 SOURCE: Byull. Eksp. Biol. Med. (1973), 75(3), 83-6
 CODEN: BEBMAE
 DOCUMENT TYPE: Journal
 LANGUAGE: Russian
 AB Swine aortic valves were submitted to the action of different reagents capable of altering their antigenic compn. and immunogenicity. Exts. from valves treated with either γ -rays or 4% formaldehyde had almost the same immunogenicity and produced the same no. of precipitin lines in agar gel diffusion tests as an ext. from fresh valves. Conditioning valve **tissues** with Na metaperiodate and **glutaraldehyde** depressed both the immunogenic and antigenic properties of xenogenic valves.

IT 111-30-8 7790-28-5
 RL: BIOL (Biological study)
 (heart valve antigen depression by)

L20 ANSWER 23 OF 23 HCAPLUS COPYRIGHT 2001 ACS
 ACCESSION NUMBER: 1966:70069 HCAPLUS
 DOCUMENT NUMBER: 64:70069
 ORIGINAL REFERENCE NO.: 64:13159d-g
 TITLE: Mechanism of a histochemical reaction differentiating between adrenaline- and noradrenaline-storing cells in the electron microscope
 AUTHOR(S): Coupland, R. E.; Hopwood, D.
 CORPORATE SOURCE: Queen's Coll., Dundee, UK
 SOURCE: Nature (1966), 209(5023), 590-1
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB cf. ibid. 201(4925), 1240-2(1964). Aq. solns. of noradrenaline (I) react with **glutaraldehyde** (II) with the production of a dense ppt. The kinetics and product of the reaction were examp. by uv and visible light and by ir spectroscopy and paper chromatography using various solvent systems; m.p. detns. were carried out and recrystn. attempted. All findings are in favor of the product being a polymer of II and I combined in a mol. ratio of 1:1. To assess the importance of II in this reaction as it relates to electron microscopy, adrenal glands were variously fixed: with Caufield's buffered (pH 7.4) osmium tetroxide-sucrose fixation, chromaffin cells contained large nos. of electron-dense chromaffin granules which exhibited a peripheral limiting membrane; using 4% II (pH 7.4) fixation only, 2 distinct types of chromaffin cells could be identified. Electron-dense granules, usually of homogeneous nature, represent the I-storing elements while the adrenaline-storing cells contain granules which have a washed-out appearance and show internal granularity. Four percent buffered II fixation followed by

post-fixation in buffered 1% osmium tetroxide or 3% K₂Cr₂O₇ showed 2 distinct types of granule visible in unstained sections and in sections stained with **permanganate** or briefly with Pb citrate. One type was intensely electron dense and homogeneous, the other had a washed-out appearance and showed internal granularity. The former was found in the cells giving a pos. color reaction for the I-II complex, the latter in others presumed to store adrenaline. In **tissues** post-fixed in osmium tetroxide, both types of granules showed a peripheral limiting membrane. The fate of adrenaline, which is apparently unaffected by II, was investigated. The majority of adrenaline is lost during II fixation and dehydration. This fact, together with the production of the I-II polymer accounts for the differential staining reaction. The staining reaction of the II-fixed I-storing cells can be obtained many months after **tissue** fixation, provided the cells are stored in water, saline or II; a delayed Cr, osmium, or iodate reaction may be performed on frozen sections of this material. Block impregnation of II-fixed adrenal with osmium tetroxide or K₂Cr₂O₇ prior to embedding in paraffin results in a good demonstration of the I islands, and this technique or the use of frozen sections is the method of choice for light microscopy. The fact that **tissues** are fixed initially in II is considered to be the most essential part of this technique.